

**Amendments to the Claims:**

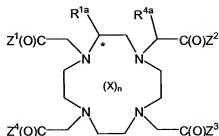
This listing of claims will replace all prior versions, and listings of claims in the application.

**Listing of Claims:**

1. (Previously presented) A method of treating a subject with cancer by administration of a macrocyclic metal chelate, said method comprising the steps of:
- (a) administering to said subject an antibody comprising an antigen recognition domain that recognizes said macrocyclic metal chelate, wherein said antibody comprises:
- a reactive site within the structure of the antibody that is not present in the wildtype of said antibody, wherein said reactive site is in a position within said antigen recognition domain and
- a targeting moiety that binds specifically to a cancer cell by binding with a member selected from a cell surface receptor and cell surface antigen, thereby forming a cell-antibody complex;
- wherein said macrocyclic metal chelate is 1,4,7,10-tetraazacyclododecane-N,N',N'',N'''-tetraacetic acid (DOTA), and comprises a reactive functional group with a reactivity complementary to said antibody reactive site; and
- (b) administering to said subject said macrocyclic metal chelate, thereby forming a covalent bond between said reactive site and said reactive functional group.

2. – 5. (Canceled).

6. (Currently amended) The method of claim 1, wherein said ~~substituted or unsubstituted DOTA~~ macrocyclic metal chelate comprises 1,4,7,10-tetraazacyclododecane-N,N',N'',N'''-tetraacetic acid (DOTA) and has the formula:



wherein

$R^{1a}$  and  $R^{4a}$  are members independently selected from H, substituted or unsubstituted alkyl, substituted or unsubstituted heteroalkyl, substituted or unsubstituted aryl and linker moieties;

X is a member selected from a lanthanide ion, an actinide ion, an alkaline earth metal ion, and a group IIIB transition metal ion;

$Z^1$ ,  $Z^2$ ,  $Z^3$  and  $Z^4$  are members independently selected from  $OR^1$  and  $NR^1R^2$

in which

$R^1$  and  $R^2$  are members independently selected from H, substituted or unsubstituted alkyl and substituted or unsubstituted heteroalkyl;

n is a member selected from 0 and 1.

7. (Cancelled).

8. (Previously presented) The method of claim 6, wherein the carbon atom marked \* is of S configuration.

9. (Cancelled)

10. (Previously presented) The method of claim 1, wherein said targeting moiety binds specifically to said cell surface antigen.

11. (Original) The method of claim 1, wherein the targeting moiety is covalently attached to said antibody.

12. (Previously presented) The method of claim 10, wherein the targeting moiety is a second antibody.

13. (Original) The method of claim 11, wherein the targeting moiety specifically binds to a protein on a cancer cell.

- 1 14. (Original) The method of claim 1, wherein the subject is a mammal.
- 1 15. (Previously presented) The method of claim 14, wherein the mammal is a human.
- 1 16. (Withdrawn) A method of *in vivo* imaging, said method comprising the steps of:  
2 (a) administering to a subject an antibody comprising an antigen recognition domain that  
3 recognizes a macrocyclic metal chelate, wherein said antibody comprises a recognition  
4 moiety that binds specifically to a cell, thereby forming a cell-antibody complex;  
5 (c) administering to said subject said metal chelate, thereby specifically binding said compound to  
6 said antibody to form a cell-antibody-metal chelate complex; and  
7 (d) detecting said cell-antibody-metal chelate complex.
- 1 17. (Withdrawn) The method of claim 16, wherein said metal chelate comprises four nitrogen atoms.
- 1 18. (Withdrawn) The method of claim 16, wherein the step of detecting is by positron emission  
2 tomography.
- 1 19. (Withdrawn) The method of claim 16, wherein the step of detecting is by magnetic +resonance  
2 imaging.
- 1 20. (Withdrawn) The method of claim 16, wherein the step of detecting is by detection of lanthanide  
2 luminescence.
- 1 21. (Withdrawn) The method of claim 16, further comprising, between steps (a) and (b),  
2 administering a clearing agent to said subject.
- 1 22. (Withdrawn) The method of claim 16, wherein the subject is a mammal.
- 1 23. (Withdrawn) The method of claim 22, wherein the mammal is a human.
- 1 24. (Previously presented) The method according to claim 1 wherein said antibody has the structure:  
2  $(Ab)_n-L-T$   
3 wherein,  
4  $n'$  is an integer selected from 1 to 10 ;  
5 Ab represents said antibody;  
6 L is a member selected from a chemical bond and a linking group that may contain one or  
7 more functional groups; and

8                   . T is said targeting moiety.

1   25.       (Canceled).

1   26.       (Previously presented) The method of claim 24, wherein said targeting moiety is a second  
2 antibody that binds specifically to a cell surface antigen.

1   27.       (Previously presented) The method according to claim 24 wherein said antibody is administered  
2 to said subject as a pharmaceutical composition comprising said antibody and a pharmaceutically  
3 acceptable carrier.

1   28.       (Canceled)

1   29.       (Cancelled).

1   30.       (Previously presented) The method according to claim 1, wherein said cell is a cancer /\*-cell.

1   31       (Canceled)

2   32.       (Cancelled).

1   33.       (Previously presented) The method according to claim 6, wherein

2               R<sup>1a</sup> and R<sup>4a</sup> are H;

3               Z<sup>1</sup>, Z<sup>2</sup>, Z<sup>3</sup> and Z<sup>4</sup> are OH;

4               and n is 1.

1   34.       (Previously presented) The method according to claim 33, wherein said targeting moiety is a  
2 second antibody that binds specifically to a cell surface antigen.

1   35.       (Previously presented) The method according to claim 34, wherein said targeting moiety is anti-  
2 CEA.

1   36.       (Previously presented) The method according to claim 33, wherein said targeting moiety is anti-  
2 CEA.

1   37.       (Currently amended) The method according to claim 1, wherein said antibody has a first  
2 sequence having at least 95 ~~percent homology~~ % sequence identity with SEQ ID NO. 1; and wherein said  
3 antibody has a second sequence having at least 95 ~~percent homology~~ % sequence identity with SEQ ID  
4 NO. 5.

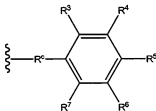
38. (Previously presented) The method according to claim 1, wherein said reactive site comprises sulfur.

39. (Cancelled) The method according to claim 1, wherein said antibody is purified.

40. (New) The method according to claim 6, wherein  $R^{1a}$  is a member independently selected from substituted or unsubstituted alkyl, substituted or unsubstituted heteroalkyl, substituted or unsubstituted aryl and linker moieties.

41. (New) The method according to claim 6, wherein  $R^{4a}$  is a member independently selected from substituted or unsubstituted alkyl, substituted or unsubstituted heteroalkyl, substituted or unsubstituted aryl and linker moieties.

42. (New) The method according to claim 6, wherein said DOTA further comprises an arylalkyl moiety having a structure according to the formula:



wherein

$R^5$  is an unsubstituted unbranched alkyl linker;

$R^3$ ,  $R^4$ ,  $R^5$ ,  $R^6$  and  $R^7$  are members independently selected from H, halogen,  $NO_2$ , CN,  $X^1R^8$ ,  $NR^9R^{10}$ , and  $C(X^2)R^{11}$ ,

wherein

$X^1$  is a member selected from O, NH, and S;

$X^2$  is a member selected from O, S, and NH;

$R^8$  and  $R^9$  are members independently selected from H, substituted or unsubstituted alkyl, substituted or unsubstituted heteroalkyl, substituted or unsubstituted heteroalkyl and  $C(Z^3)R^{12}$

wherein

$Z^3$  is a member selected from O, S and NH;

$R^{12}$  is a member selected from substituted or unsubstituted alkyl, substituted or unsubstituted heteroalkyl and  $OR^{13}$

wherein

19                               R<sup>13</sup> is a member selected from substituted or unsubstituted alkyl,  
20                               substituted or unsubstituted heteroalkyl, substituted or  
21                               unsubstituted aryl, and substituted or unsubstituted heteroaryl;  
22               R<sup>10</sup> is a member selected from H, substituted or unsubstituted alkyl, substituted or  
23               unsubstituted heteroalkyl, and OH, and  
24       R<sup>9</sup> and R<sup>10</sup> taken together are optionally (=C=S);  
25               R<sup>11</sup> is a member selected from H, halogen, substituted or unsubstituted alkyl, substituted  
26               or unsubstituted heteroalkyl, OR<sup>14</sup>, and NR<sup>15</sup>R<sup>16</sup>,  
27       wherein  
28               R<sup>14</sup> is a member selected from H, substituted or unsubstituted alkyl, substituted  
29               or unsubstituted heteroalkyl, and C(O)R<sup>17</sup>,  
30       wherein  
31               R<sup>17</sup> is a member selected from substituted or unsubstituted alkyl, and  
32               substituted or unsubstituted heteroalkyl; and  
33       R<sup>15</sup> and R<sup>16</sup> are members independently selected from H, substituted or  
34               unsubstituted alkyl, and substituted or unsubstituted heteroalkyl.